

FIG. 1

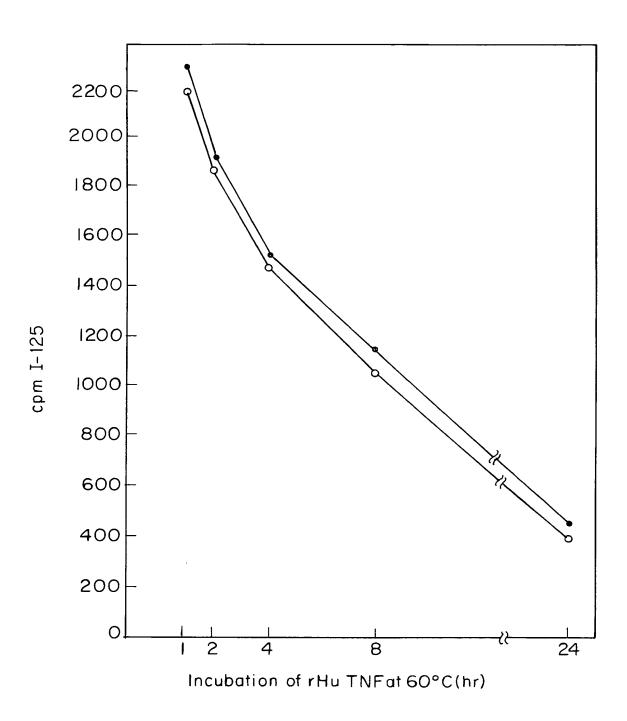
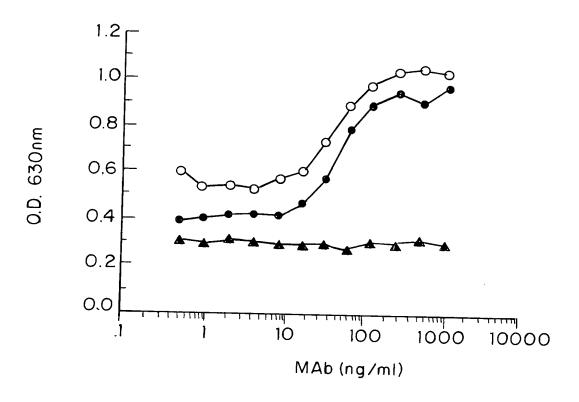


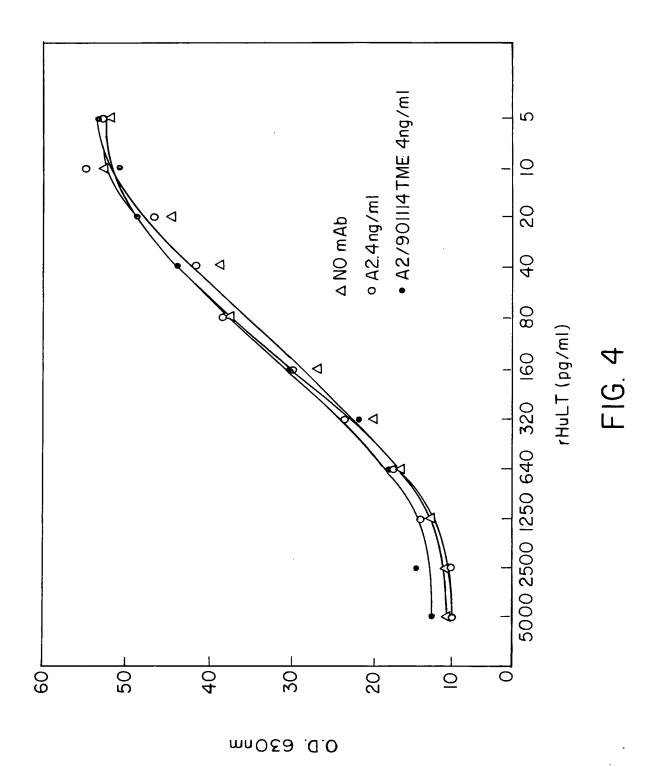
FIG. 2

Title: Methods of Treating Neurodegenerative........
Inventors: Junming Le, et al.

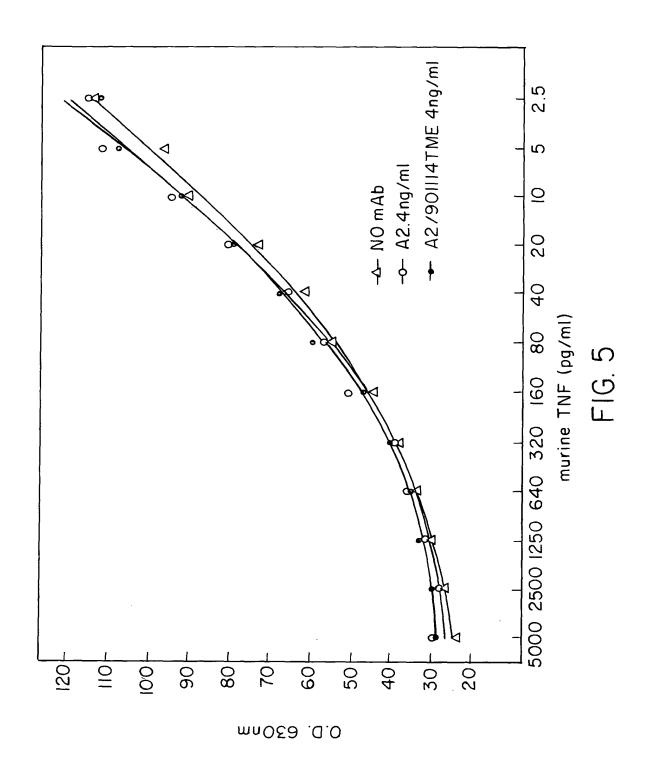


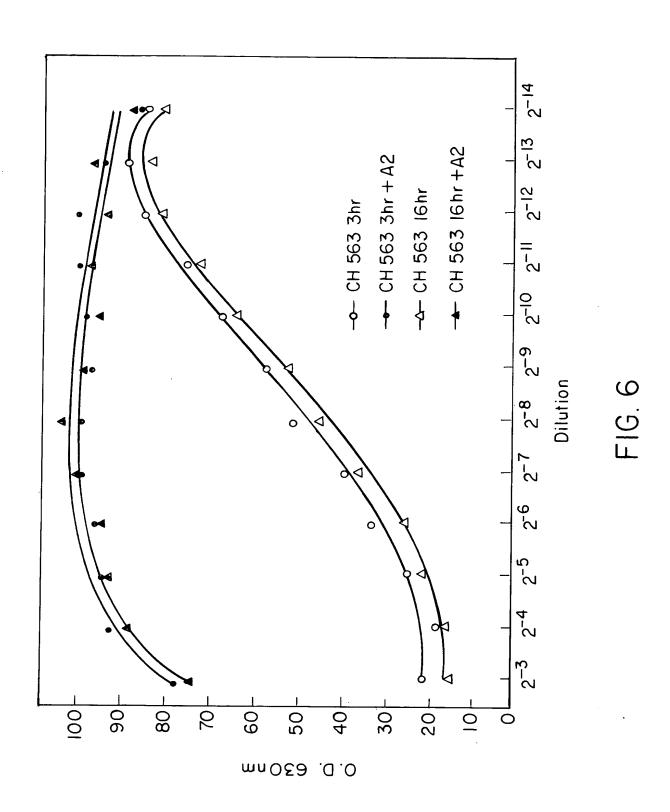
- Natural Human TNF
- Recombinant Human TNF
- ▲ Control

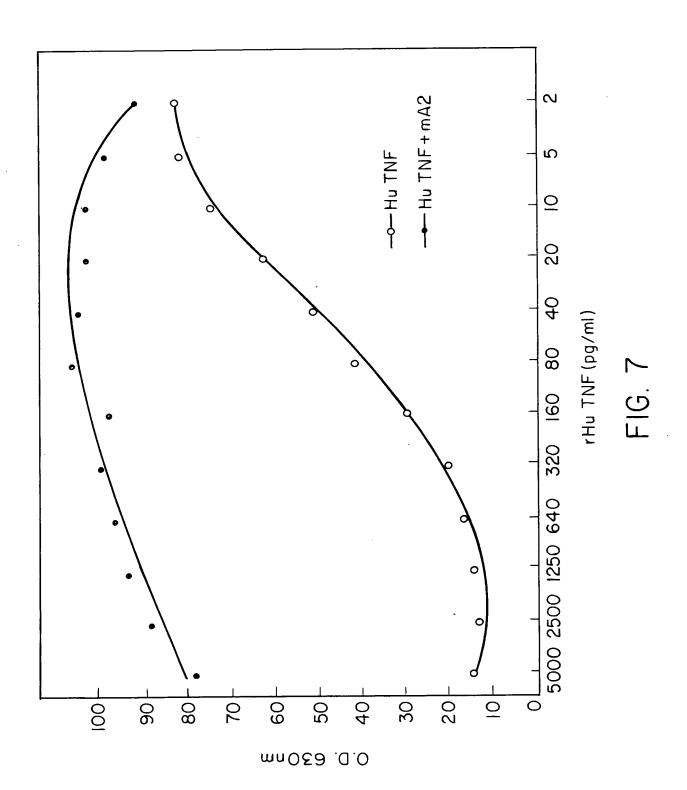
FIG. 3











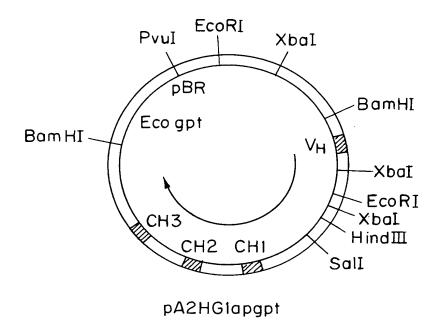


FIG. 8A

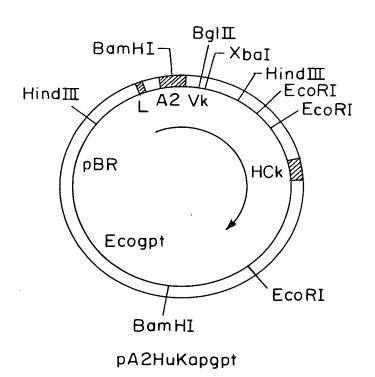


FIG. 8B

Title: Methods of Treating Neurodegenerative..........
Inventors: Junming Le, et al.

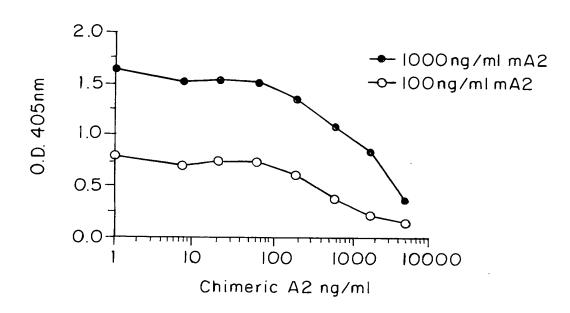


FIG. 9A

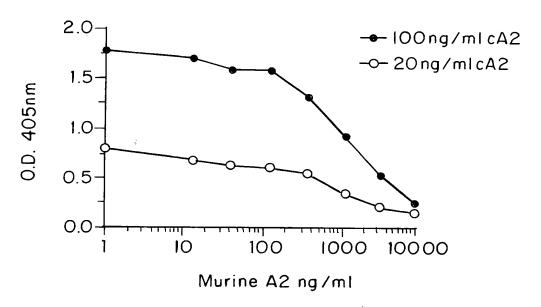


FIG. 9B

Docket No. 0975.1005-036

Title: Methods of Treating Neurodegenerative.......

Inventors: Junming Le, et al.

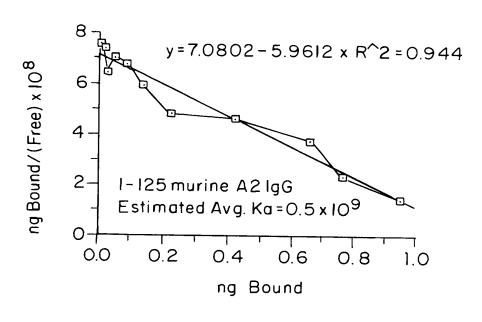


FIG. 10A

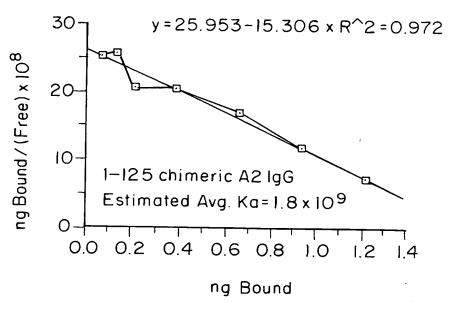


FIG. 10B

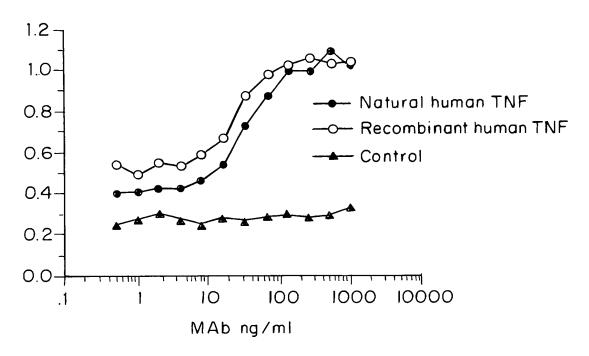


FIG. 11

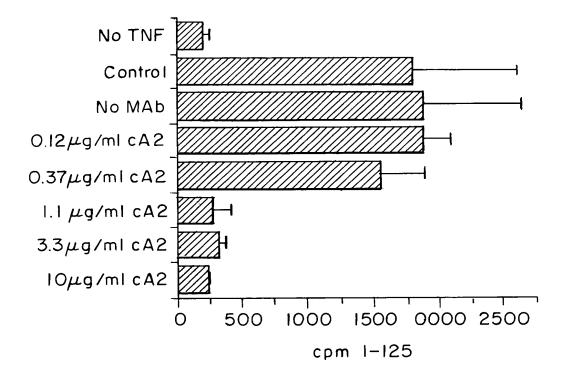


FIG. 12

	Asp	Leu	Pro	Ile	Ser	Gly	Pro
	Pro	Tyr	Ser	Thr	Tyr	Asn	
	Arg	Ile			Ile	Ala	Ala
Leu	Asn		Ile Lys	Thr His	Leu	Leu	Val
Ala	Ile Asn	Glu	Ala		${ m Tyr}$	Leu	Val
Ile	Glu	$\mathrm{T}\mathrm{y}\mathrm{r}$	Ser	Leu	ren	Ala	His
Ile	Ala	Trp	Leu	Val	$_{ m G1y}$	Asn	Ala
Gly	Ser	Pro	Leu	His	Glu	Ala	Val
Phe	Leu	Lys	Asn	Thr	Ser	Arg	Pro Val Ala His Val Val Ala Asn
Tyr	130 Asp Arg Leu Ser Ala Glu	Ala Lys Pro Trp Tyr Glu Pro	Val Asn Leu Leu Ser Ala	70 Pro Ser Thr His Val Leu Leu	Pro	Arg	Lys
150 Glu Ser Gly Gln Val Tyr Phe Gly Ile Ile Ala	130 Asp	110 Glu	90 Lys	70 Pro	50 Val Pro Ser Glu Gly Leu Tyr Leu Ile	30 Leu Gln Trp Leu Asn Arg Arg Ala Asn Ala Leu Leu Ala Asn Gly	10 Arg Thr Pro Ser Asp Lys
Gln	G1y		${ m Th} r$	Cys	Asn Gln Leu Val	Leu	Ser
Gly	Lys Gly	Gly Ala	Ser Tyr Gln Thr	Gly Gln Gly Cys	Leu	Trp	Pro
Ser	Leu Glu	Pro Glu	$\mathrm{T} \mathrm{yr}$	Gln	Gln	Gln	\mathtt{Thr}
		Pro	Ser	$_{ m G1y}$	Asn	Leu	Arg
Phe Ala	Gln	\mathtt{Thr}	Val	Lys	Asp	Gln	Ser
Phe	Phe	Glu	Arg Ile Ala	Val Leu Phe	Glu Leu Arg Asp	Ala Glu Gly Gln	Ser
Leu Asp	Val	Gln Arg	Ile	Leu	Leu	Glu	Ser
	. G1y						Arg
141 Tvr	121 Gly	101 Cys	81 Ser	61 Gln	41 Val	21 Gln	1 Val

FIG. 1

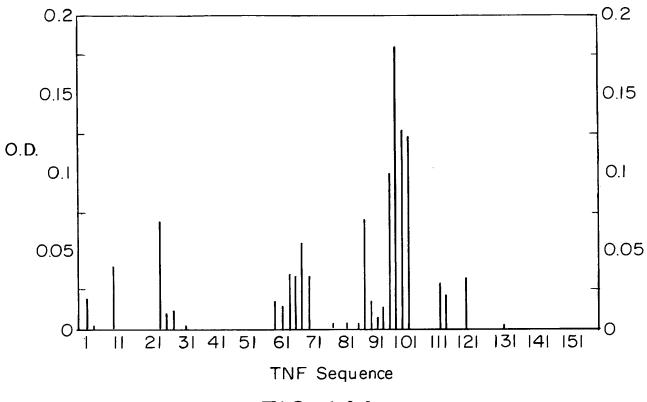


FIG. 14A

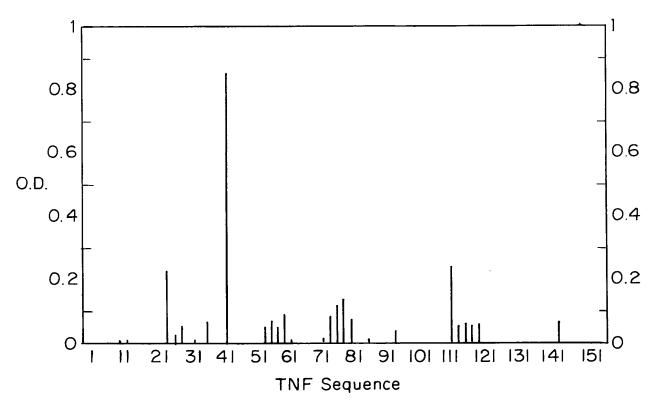


FIG. 14B

	Pro
	Asn
	Ala
	Val A
	1.
	Va
	His Val
	Ala
	Val Ala
•	Pro
	Lys
10	Asp
	Ser
	Pro
	Thr
	Arg
	Ser
	Ser
	Ser
	Arg :
\vdash	Val /

	31y
	•
	a Asn
	Leu Ala
	Leu
	a Leu L
	Ala I
	n A
•	Asn
	Ala
	g Arg A
	Arg .
30	Asn
	Leu
	\mathtt{Trp}
	31n
	_
	Leu
	Gln
	Gly
	glu (
	Ala
21	Gln

50	l Glu Leu Arg Asp Asn Gln Leu Val Val Pro Ser Glu Gly Leu Tyr Leu Ile Tyr Ser	
	g As	
	u Arç	
	l Leι	
	l G11	
4 T	Val	

Ile
Thr
His
Thr I
I na
Leu Leu
Val
His
Thr
Ser
70 Pro S
Gly Cys
Gln
Gly
Lys
Phe
Leu]
Val I
l ln V
<u>6</u>

Pro	Leu
Ser Pro	Tyr
Lys	Ile
Ile	Pro
: Ala Ile Lys	Glu
Ser	Tyr
Leu	Trp
Leu	Pro
90 Gln Thr Lys Val Asn Leu Leu Ser	Gly Ala Glu Ala Lys Pro Trp Tyr Glu Pro Ile Tyr Leu
Val	Ala
90 Lys	110 Glu
Thr	Ala
Gln	Gly
77	Glu
Ser T	Pro
Val	Thr
Ala	Glu
Ile	Arg Glu Thr P
Arg	Gln
81 Ser	101 Cys

	Asp
	Pro
	Arg
	Asn
	Ile
	Glu
	Ala
	Ser
	Asp Arg Leu
130	Asp
	Gly
	Lys
	Glu
	Leu
	Gln
	Phe
	Val
	Gly ,
121	Gly (

	Leu
	Ala
	I1e
	, Ile
	Gly
	Phe
	Tyr
150	Val
	Gln
	G1y
	Ser
	Glu
	Ala
	Phe
	Asp
	Leu
141	Tyr

Docket No. 0975.1005-030 Title: Methods of Treating Neurodegenerative....... Inventors: Junming Le, *et al*.

GACATCTTGCTGACTCAGTCTCCAGCCATCCTGTCTGTGAGTCCAGGAGAAAGAGTCCAGT AspileLeuLeuThrGlnSerProAlaIleLeuSerValSerProGlyGluArgValSer TTCTCCTGCAGGCCAGTCAGTTCGTTGGCTCAAGCATCCACTGGTATCAGCAAGAACA $\tt Phe SerCysArgAlaSerGlnPhe ValGlySerSerIleHisTrpTyrGlnGlnArgThr$

AATGGTTCTCCAAGGCTTCTCATAAAGTATGCTTCTGAGTCTATGTCTGGGATCCCTTCC ${\tt AsnGlySerProArgLeuLeuIleLysTyrAlaSerGluSerMetSerGlyIleProSer}$

AGGTTTAGTGGCAGTGGATCAGGGACAGATTTTACTCTTAGCATCAACACTGTGGAGTCT ${\tt ArgPheSerGlySerGlySerGlyThrAspPheThrLeuSerIleAsnThrValGluSer}$

GAAGATATTGCAGATTATTACTGTCAAGAAAGTCATAGCTGGCCATTCACGTTCGGCTCG GluAspIleAlaAspTyrTyrCysGlnGlnSerHisSerTrpProPheThrPheGlySer

GGGACAAATTTGGAAGTAAAA GlyThrAsnLeuGluValLys FIG. 16A

Inventors: Junming Le, et al.

GAAGTGAAGCTTGAGGAGTCTGGAGGAGGCTTGGTGCAACCTGGAGGATCCATGAAACTC ${\tt GluValLysLeuGluGluSerGlyGlyGlyLeuValGlnProGlyGlySerMetLysLeu}$

SerCysValAlaSerGlyPheIlePheSerAsnHisTrpMetAsnTrpValArgGlnSer TCCTGTGTTGCCTCTGGATTCATTTTCAGTAACCACTGGATGAACTGGGTCCGCCAGTCT

CCAGAGAAGGGGCTTGAGTTGCTTGAAATTAGATCAAAATCTATTAATTCTGCAACA $\tt ProGluLysGlyLeuGluTrpValAlaGluIleArgSerLysSerIleAsnSerAlaThr$

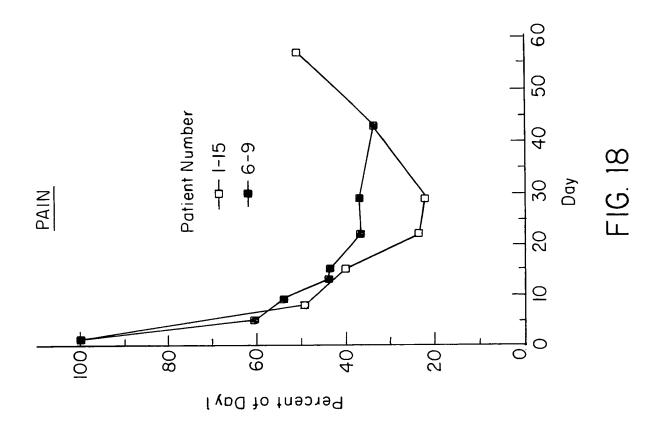
HisTyrAlaGluSerValLysGlyArgPheThrIleSerArgAspAspSerLysSerAla CATTATGCGGAGTCTGTGAAAGGGAGGTTCACCATCTCAAGAGAGATGATTCCAAAAGTGCT

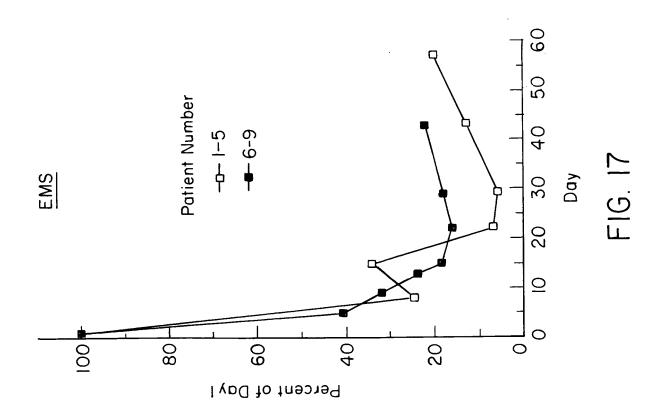
GTGTACCTGCAAATGACCGACTTAAGAACTGAAGACACTGGCGTTTATTACTGTTCCAGG $\verb|ValTyrLeuGlnMetThrAspLeuArgThrGluAspThrGlyValTyrTyrCysSerArg| \\$

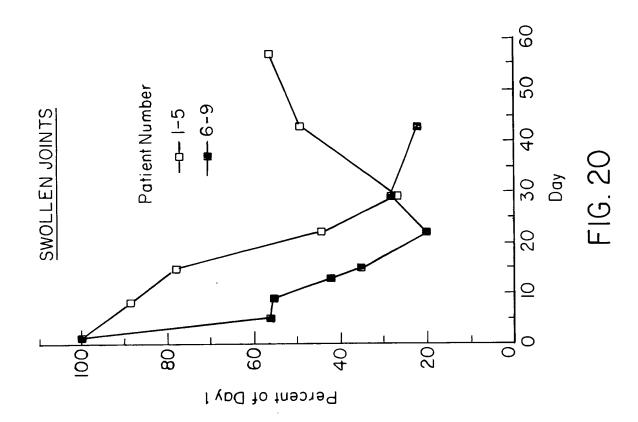
AsnTyrTyrGlySerThrTyrAspTyrTrpGlyGlnGlyThrThrLeuThrValSer AATTACTACGGTAGTACCTACGACTACTGGGGCCAAGGCACCACTCTCACAGTGTCC

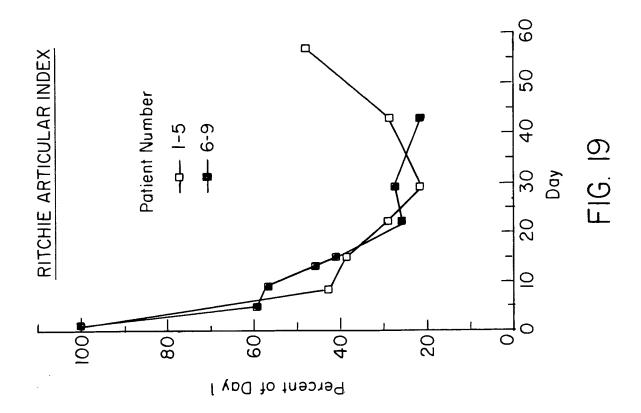
FIG. 16B

Docket No. 0975.1005-036 Title: Methods of Treating Neurodegenerative...... Inventors: Junming Le, *et al*.



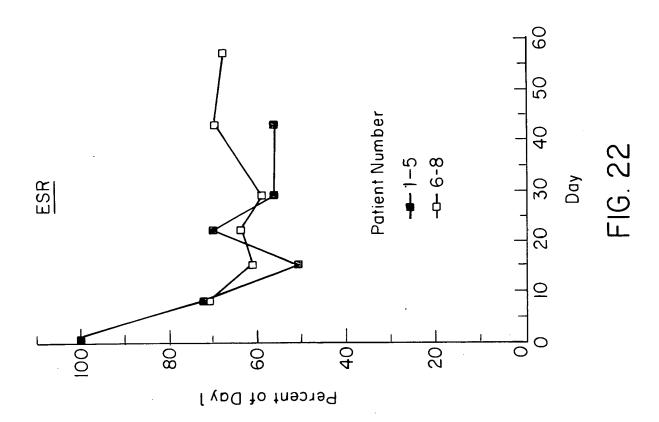


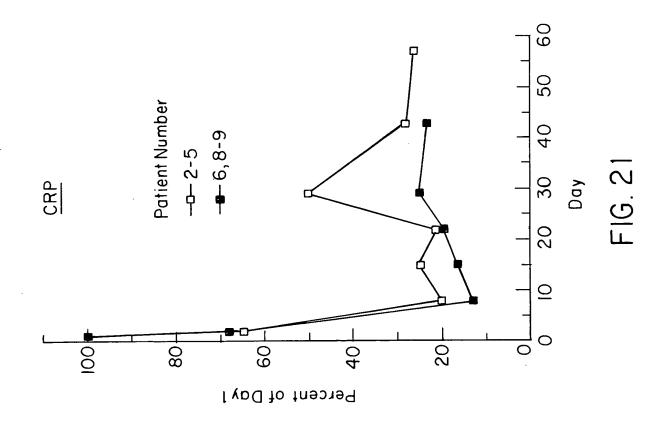




Docket No. 0975.1005-036

Title: Methods of Treating Neurodegenerative....... Inventors: Junming Le, *et al.*





Docket No. 0975.1005-036 Title: Methods of Treating Neurodegenerative....... Inventors: Junming Le, *et al.*

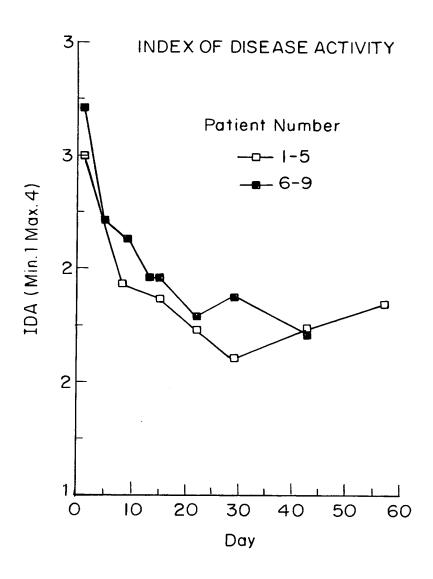
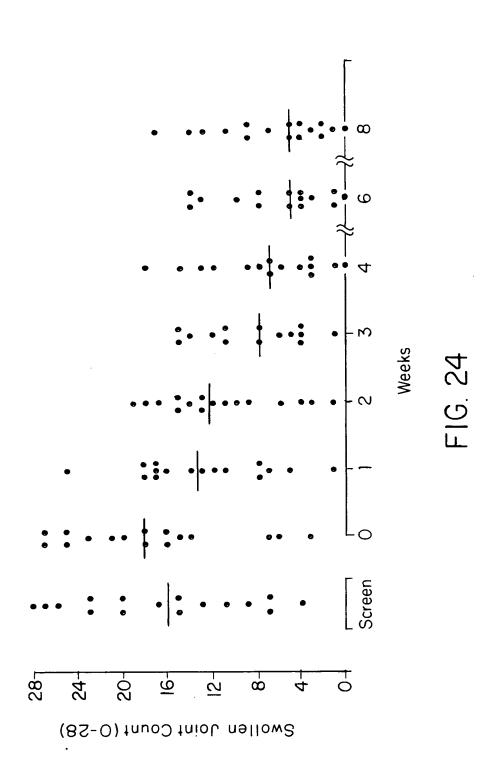


FIG. 23

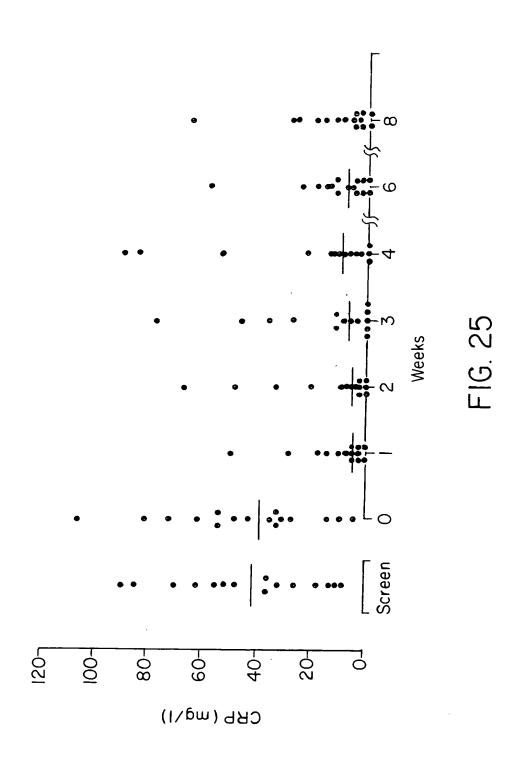
Docket No. 0975.1005-036

Title: Methods of Treating Neurodegenerative......



Docket No. 0975.1005-036

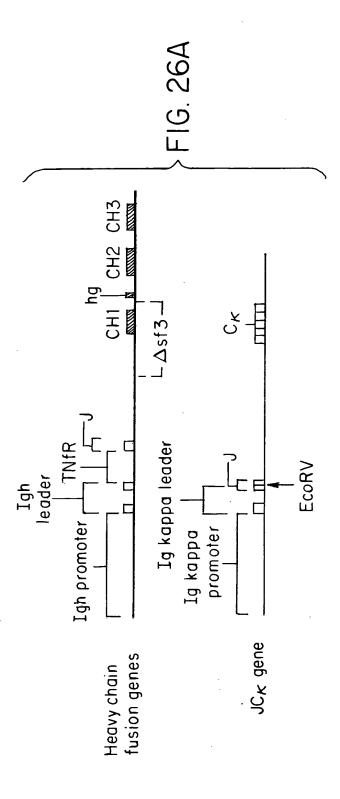
Title: Methods of Treating Neurodegenerative......

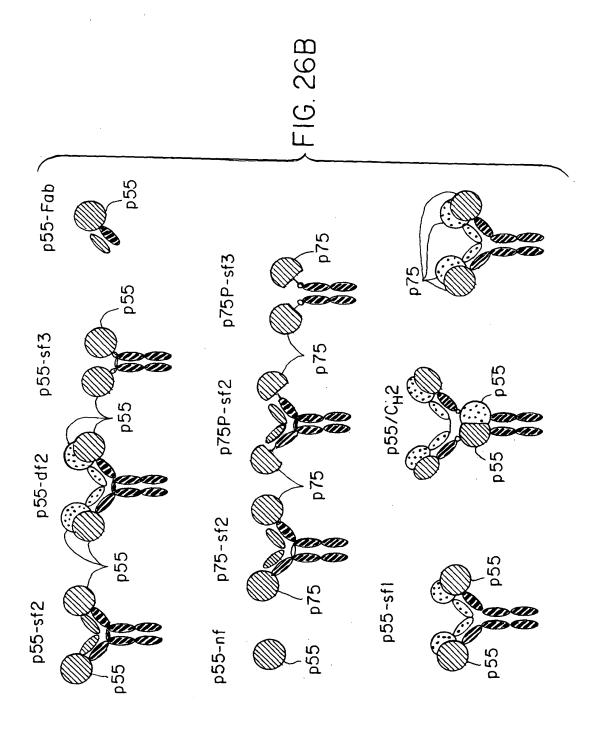


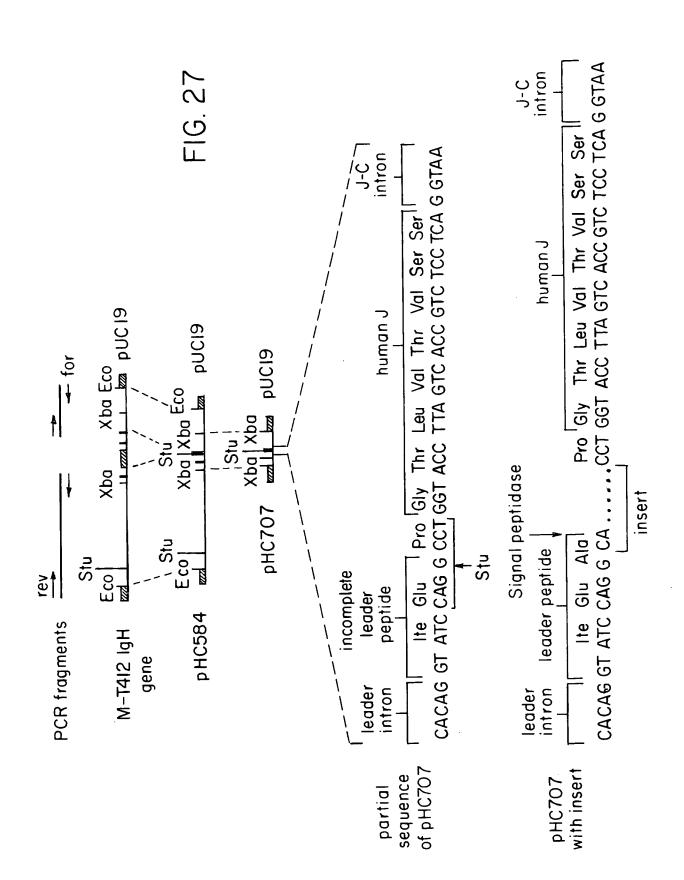
Docket No. 0975.1005-036

Title: Methods of Treating Neurodegenerative......

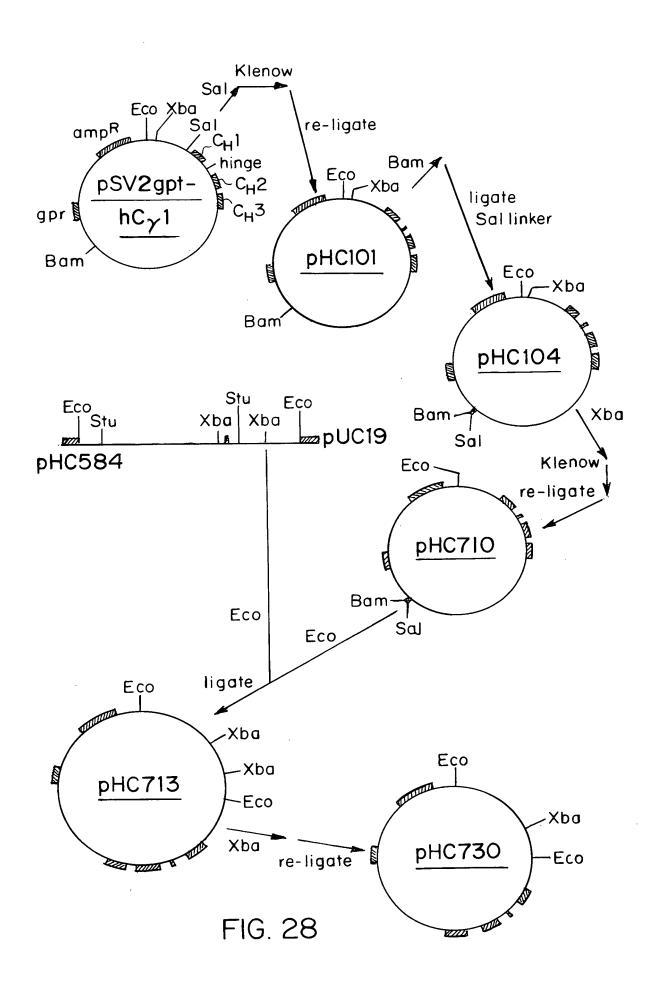
Inventors: Junming Le, et al.

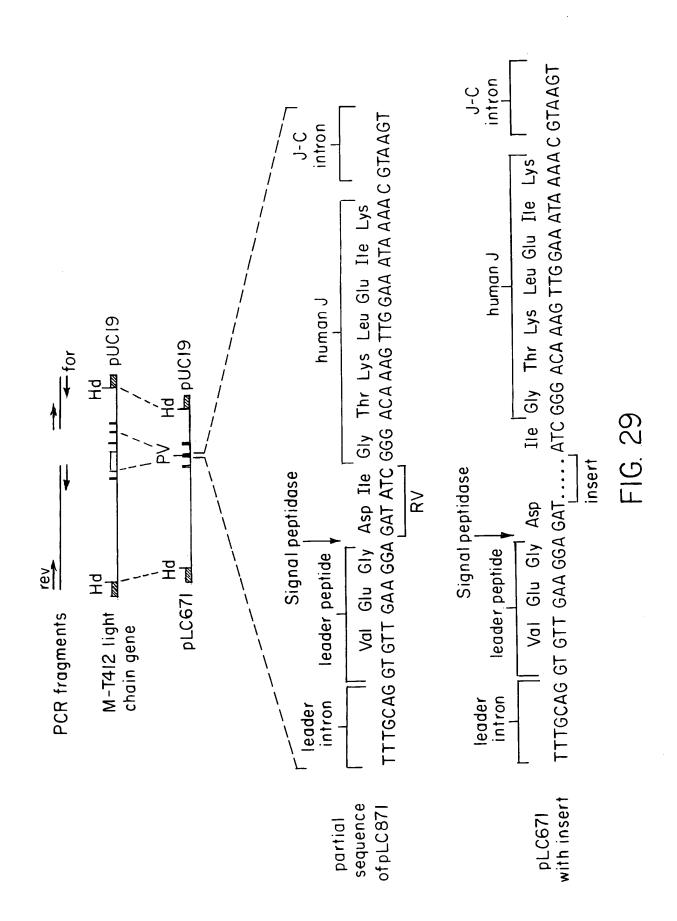






Docket No. 0975.1005-036 Title: Methods of Treating Neurodegenerative....... Inventors: Junming Le, *et al.*





Docket No. 0975.1005-030 Title: Methods of Treating Neurodegenerative....... Inventors: Junming Le, et al.

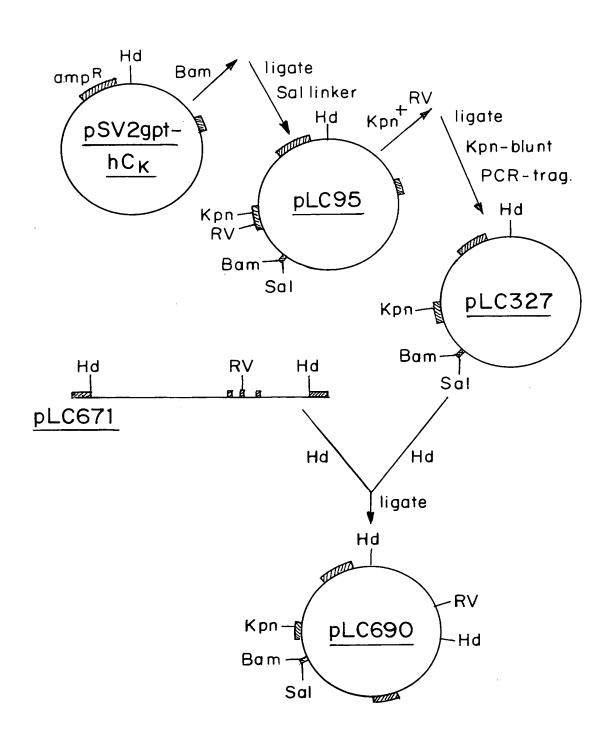


FIG. 30

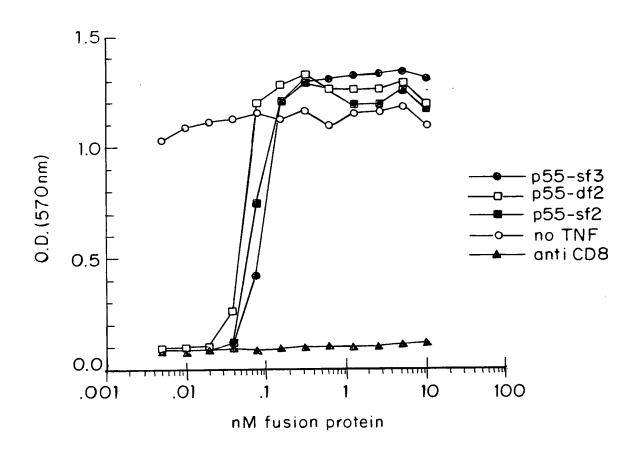


FIG. 31A

Docket No. 0975.1005-030 Title: Methods of Treating Neurodegenerative....... Inventors: Junming Le, *et al.*

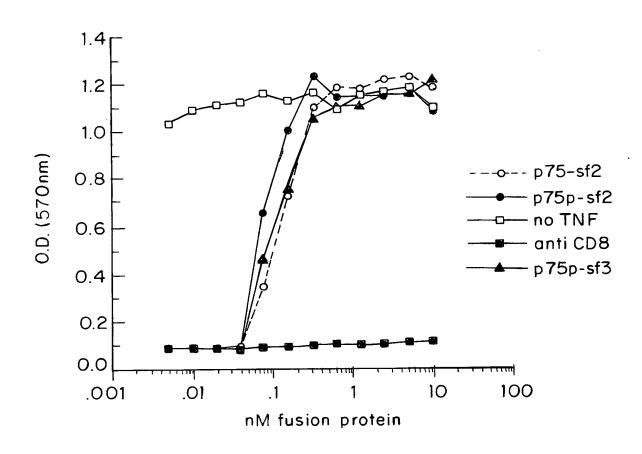


FIG. 31B

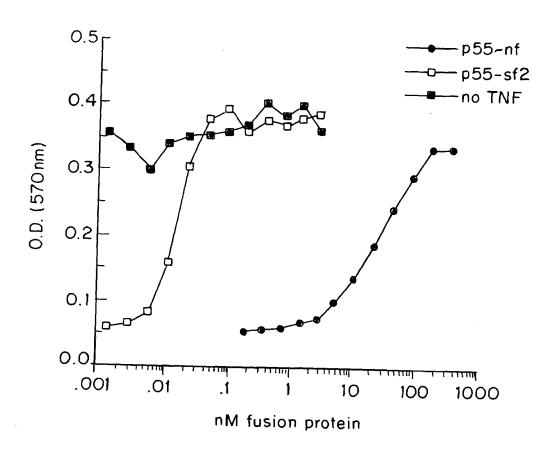


FIG. 31C

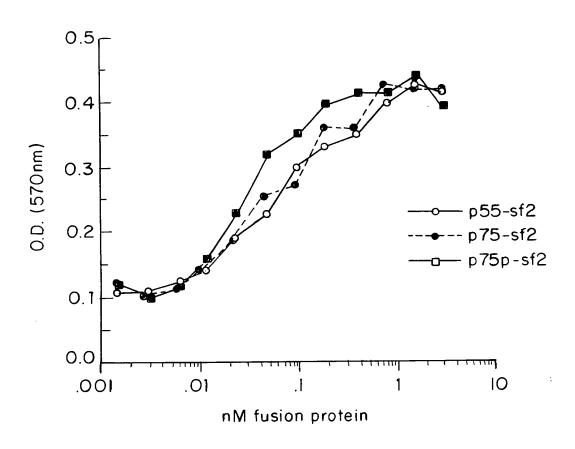


FIG. 32

Docket No. 0975.1005-030 Title: Methods of Treating Neurodegenerative....... Inventors: Junming Le, *et al.*

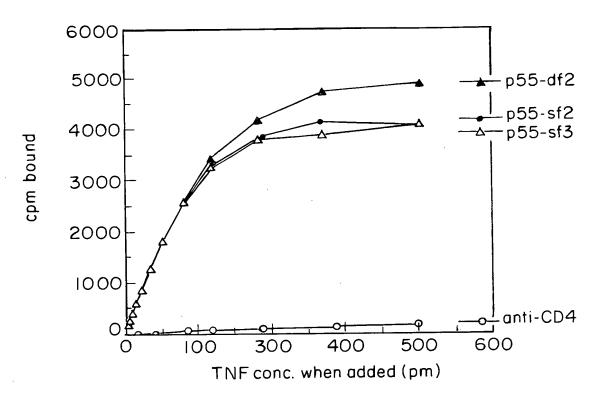


FIG. 33A

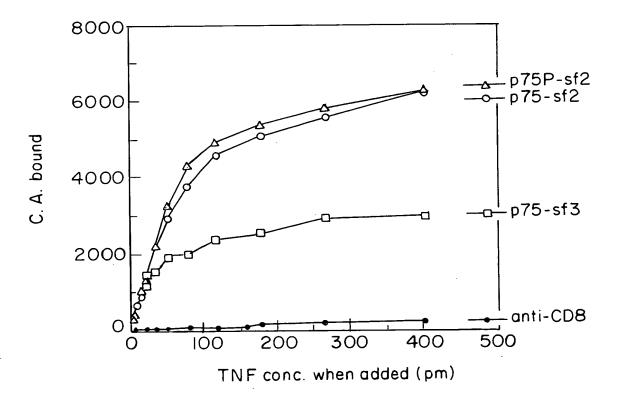
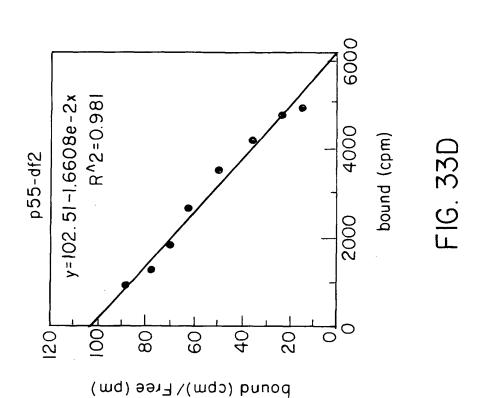
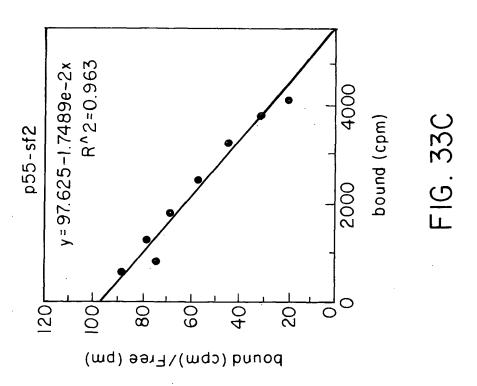
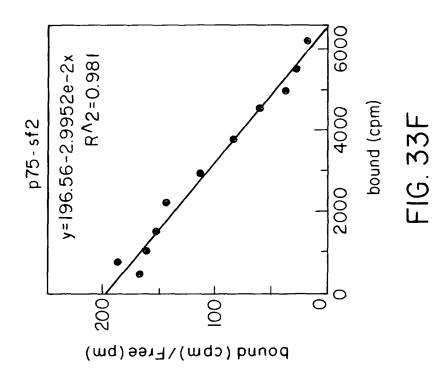


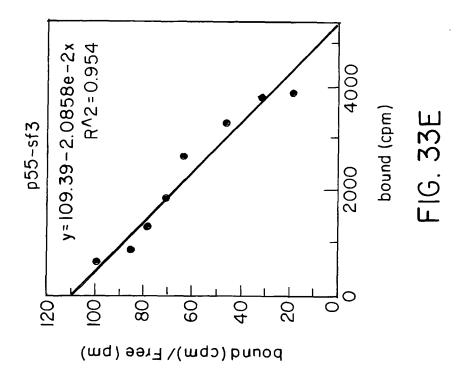
FIG. 33B



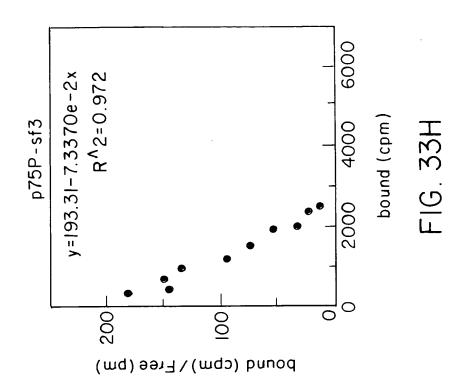


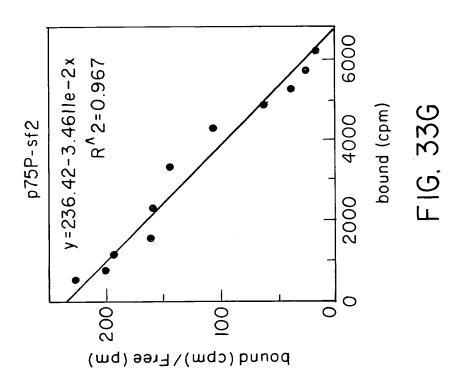
Docket No. 0975.1005-036 Title: Methods of Treating Neurodegenerative....... Inventors: Junming Le, *et al.*

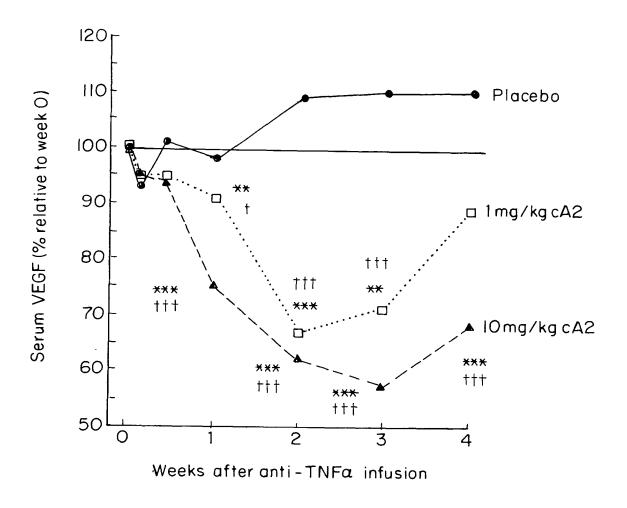




Docket No. 0975.1005-036 Title: Methods of Treating Neurodegenerative....... Inventors: Junming Le, *et al.*







* p \leq 0.05, **p \leq 0.01, *** p \leq 0.001 *versus* pre-infusion † p \leq 0.05, ††p \leq 0.01, ††† p \leq 0.001 *versus* change in placebo group

FIG. 34